

Inventor(s): BOUCHARD *et al.*
Application No.: 08/786,937
Attorney Docket No.: 098501-0235299

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I. AMENDMENTS TO THE CLAIMS

1-37. (Canceled)

38. (Currently Amended) A method for obtaining the production of a fertilizable oocyte within a program of controlled ovarian stimulation for assisted reproduction techniques (COS/ART) comprising:

- (a) administering an exogenous gonadotropin to induce follicle growth, and
- (b) administering a luteinizing hormone releasing hormone (LHRH) antagonist to prevent a premature LH surge, wherein the LHRH antagonist is administered in a single or dual dosage regimen of 1 to 10 mg per dose beginning on menstruation cycle day 1 to 10; and wherein follicular growth occurs in the absence of a LH surge, a fertilizable oocyte is produced, ovulation occurs between day 9 and 20 of the menstruation cycle, and the LHRH antagonist is sufficient to suppress LH, while FSH secretion is maintained at a natural level and individual estrogen development is not affected.

39. (Previously Presented) The method of claim 38, wherein the dosage of LHRH antagonist is in the range of 2-6 mg per dose.

40. (Previously Presented) The method of claim 38, wherein dosage of LHRH antagonist is 3 mg per dose.

41. (Cancel)

42. (Previously Presented) The method of claim 38, wherein the LHRH antagonist is administered by subcutaneous injection.

43. (Cancel)

44. (Previously Presented) The method of claim 38, wherein the LHRH antagonist is administered starting cycle day 4 to 8.

45. (Previously Presented) The method of claim 38, wherein the LHRH antagonist is administered starting on cycle day 6 to 10 and ovulation occurs between day 9-16 of the menstruation cycle.

Inventor(s): BOUCHARD *et al.*
Application No.: 08/786,937
Attorney Docket No.: 098501-0235299

46. (Previously Presented) The method of claim 38, wherein ovulation occurs within 6.5 days following administration of a single or second dose of the LHRH antagonist.

47. (Previously Presented) The method of claim 38, wherein ovulation occurs normally, without the administration of a hormone or hormone agonist to induce ovulation.

48. (Previously Presented) The method of claim 38, wherein ovulation is induced by administering a hormone or hormone agonist in order to induce ovulation.

49. (Previously Presented) The method of claim 38, wherein ovulation is induced by administering a hormone or hormone agonist selected from the group consisting of native LH, recombinant LH, an LHRH agonist, and HCG.

50. (Previously Presented) The method of claim 38, wherein the LHRH antagonist is Cetrorelix.

51. (Currently Amended) A method for obtaining the production of a fertilizable oocyte within a program of COS/ART comprising:

(a) administering human menopausal gonadotropin (HMG) to induce follicle growth, and

(b) administering Cetrorelix to prevent a premature LH surge, wherein Cetrorelix is administered in a single or dual dosage regimen of 1 to 10 mg per dose beginning on menstruation cycle day 1 to 10; and

whereby wherein follicular growth occurs in the absence of a LH surge, a fertilizable oocyte is produced, ovulation occurs between day 9 and 20 of the menstruation cycle, and the Cetrorelix is sufficient to suppress LH, while FSH secretion is maintained at a natural level and individual estrogen development is not affected.

52. (Previously Presented) The method of claim 51, wherein the dosage of the LHRH antagonist is in the range of 2-6 mg per dose.

53. (Previously Presented) The method of claim 51, wherein the dosage of Cetrorelix is 3 mg per dose.

54. (Cancel)

55. (Cancel)

Inventor(s): BOUCHARD *et al.*
Application No.: 08/786,937
Attorney Docket No.: 098501-0235299

56. (Previously Presented) The method of claim 51, wherein the Cetorelix is administered starting cycle day 4 to 8.

57. (Previously Presented) The method of claim 51, wherein Cetorelix is administered starting on cycle day 6 to 10 and ovulation occurs between day 9-16 of the menstruation cycle.

58. (Previously Presented) The method of claim 51, wherein ovulation occurs within 6.5 days following administration of a single or second dose of Cetorelix.

59. (Previously Presented) The method of claim 51, wherein ovulation occurs normally, without the administration of a hormone or hormone agonist to induce ovulation.

60. (Previously Presented) The method of claim 51, wherein ovulation is induced by administering a hormone or hormone agonist selected from the group consisting of native LH, recombinant LH, an LHRH agonist, and HCG.

61. (Currently Amended) An improved method for obtaining the production of a fertilizable oocyte within a program of COS/ART comprising:

- (a) administering an exogenous gonadotropin to induce follicle growth; and
- (b) administering an LHRH antagonist to prevent a premature LH surge;

wherein the improvement comprises administering the LHRH antagonist in a single or dual dosage regimen of 1 to 10 mg per dose beginning on menstruation cycle day 1 to 10, and wherein the follicular growth occurs in the absence of a LH surge, a fertilizable oocyte is produced, ovulation occurs between day 9 and 20 of the menstruation cycle, and the LHRH antagonist is sufficient to suppress LH, while FSH secretion is maintained at a natural level and individual estrogen development is not affected.

62. (Previously Presented) The improved method of claim 61, wherein the dosage of LHRH antagonist is in the range of 2-6 mg per dose.

63. (Previously Presented) The improved method of claim 61, wherein the dosage of LHRH antagonist is 3 mg per dose.

64. (Cancel)

65. (Previously Presented) The improved method of claim 61, wherein the LHRH antagonist is administered by subcutaneous injection.

Inventor(s): BOUCHARD *et al.*
Application No.: 08/786,937
Attorney Docket No.: 098501-0235299

66. (Cancel)

67. (Previously Presented) The improved method of claim 61, wherein the LHRH antagonist is administered starting cycle day 4 to 8.

68. (Previously Presented) The improved method of claim 61, wherein the LHRH antagonist is administered starting on cycle day 6 to 10 and ovulation occurs between day 9-16 of the menstruation cycle.

69. (Previously Presented) The improved method of claim 61, wherein ovulation occurs within 6.5 days following administration of a single or second dose of the LHRH antagonist.

70. (Previously Presented) The improved method of claim 61, wherein ovulation occurs normally, without the administration of a hormone or hormone agonist to induce ovulation.

71. (Previously Presented) The improved method of claim 61, wherein ovulation is induced by administering a hormone or hormone agonist selected from the group consisting of native LH, recombinant LH, an LHRH agonist, and HCG.

72. (Previously Presented) The improved method of claim 61, wherein the LHRH antagonist is Cetrorelix.

73. (Currently Amended) The improved method of claim 61 further comprising:

(a) administering human menopausal gonadotropin (HMG) to induce follicle growth; and

(b) administering Cetrorelix to prevent a premature LH surge;

wherein the improvement comprises subcutaneously administering Cetrorelix in a single or dual dosage regimen of 1 to 10 mg per dose beginning on menstruation cycle day 1 to 10; and

~~whereby~~ wherein ovulation occurs between day 9 and 20 of the menstruation cycle, and the LHRH antagonist is sufficient to suppress LH, while FSH secretion is maintained at a natural level and individual estrogen development is not affected.

74. (Previously Presented) The improved method of claim 73, wherein the dosage of Cetrorelix is in the range of 2-6 mg per dose.

Inventor(s): BOUCHARD *et al.*
Application No.: 08/786,937
Attorney Docket No.: 098501-0235299

75. (Previously Presented) The improved method of claim 73, wherein the dosage of LHRH antagonist is 3 mg per dose.

76. (Cancel)

77. (Cancel)

78. (Previously Presented) The improved method of claim 73, wherein the LHRH antagonist is administered starting cycle day 4 to 8.

79. (Previously Presented) The improved method of claim 73, wherein Cetorelix is administered starting on cycle day 6 to 10 and ovulation occurs between day 9-16 of the menstruation cycle.

80. (Previously Presented) The improved method of claim 73, wherein ovulation occurs within 6.5 days following administration of a single or second dose of Cetorelix.

81. (Previously Presented) The improved method of claim 73, wherein ovulation occurs normally, without the administration of a hormone or hormone agonist to induce ovulation.

82. (Previously Presented) The improved method of claim 73, wherein ovulation is induced by administering a hormone or hormone agonist selected from the group consisting of native LH, recombinant LH, an LHRH agonist, and HCG.

83. (Currently Amended) A method for obtaining the production of a fertilizable oocyte within a program of controlled ovarian stimulation for assisted reproduction techniques (COS/ART) comprising

(a) administering an exogenous gonadotropin to induce follicle growth,

(b) administering a luteinizing hormone releasing hormone (LHRH) antagonist to prevent a premature LH surge, wherein the LHRH antagonist is administered in a dosage regimen of daily doses of 0.25 mg/day for multiple days,

wherein the LHRH antagonist is administered daily beginning on menstruation cycle day 1 to 10, wherein the follicular growth occurs in the absence of a LH surge, a fertilizable oocyte is produced, ovulation occurs between day 9 and 20 of the menstruation cycle, and the LHRH antagonist is sufficient to suppress LH, while FSH secretion is maintained at a natural level and individual estrogen development is not affected.

Inventor(s): BOUCHARD *et al.*

Application No.: 08/786,937

Attorney Docket No.: 098501-0235299

84. (Previously Presented) The method of claim 83, wherein the LHRH antagonist is administered by subcutaneous injection.

85. (Canceled)

86. (Previously Presented) The method of claim 83, wherein the LHRH antagonist is administered starting cycle day 4 to 8.

87. (Previously Presented) The method of claim 83, wherein a daily dose of the LHRH antagonist is administered for 3 to 14 days.

88. (Previously Presented) The method of claim 83, wherein a daily dose of the LHRH antagonist is administered for 3 to 7 days.

89. (Previously Presented) The method of claim 83, wherein ovulation occurs normally, without the administration of a hormone or hormone agonist to induce ovulation.

90. (Previously Presented) The method of claim 83, wherein ovulation is induced by administering a hormone or hormone agonist selected from the group consisting of native LH, recombinant LH, an LHRH agonist, and HCG.

91. (Previously Presented) The method of claim 83, wherein the LHRH antagonist is Cetrorelix.

92. (Currently Amended) A method for obtaining the production of a fertilizable oocyte within a program of COS/ART comprising:

(a) administering human menopausal gonadotropin (HMG) to induce follicle growth, and;

(b) administering Cetrorelix to prevent a premature LH surge, wherein Cetrorelix is subcutaneously administered in a dosage regimen of daily doses of 0.25 mg per day for multiple days;

~~whereby~~ wherein follicular growth occurs in the absence of a LH surge, a fertilizable oocyte is produced, ovulation occurs between day 9 and 20 of the menstruation cycle, and the Cetrorelix is sufficient to suppress LH, while FSH secretion is maintained at a natural level and individual estrogen development is not affected.

93. (Canceled)

Inventor(s): BOUCHARD *et al.*
Application No.: 08/786,937
Attorney Docket No.: 098501-0235299

94. (Previously Presented) The method of claim 92, wherein Cetorelix is administered starting cycle day 4 to 8.

95. (Previously Presented) The method of claim 92, wherein a daily dose of Cetorelix is administered for 3 to 14 days.

96. (Previously Presented) The method of claim 92, wherein a daily dose of Cetorelix is administered for 3 to 7 days.

97. (Previously Presented) The method of claim 92, wherein ovulation occurs normally, without the administration of a hormone or hormone agonist to induce ovulation.

98. (Previously Presented) The method of claim 92, wherein ovulation is induced by administering a hormone or hormone agonist selected from the group consisting of native LH, recombinant LH, an LHRH agonist, and HCG.

99. (Previously Presented) An improved method for obtaining the production of a fertilizable oocyte within a program of controlled ovarian stimulation for assisted reproduction techniques (COS/ART) comprising

- (a) administering an exogenous gonadotropin to induce follicle growth, and
- (b) administering a luteinizing hormone releasing hormone (LHRH) antagonist to prevent a premature LH surge,

wherein the improvement comprises administering the LHRH antagonist in a dosage regimen of daily doses of 0.25 mg per day for multiple days, the follicular growth occurs in the absence of a LH surge, a fertilizable oocyte is produced, ovulation occurs between day 9 and 20 of the menstruation cycle, and the LHRH antagonist is sufficient to suppress LH, while FSH secretion is maintained at a natural level and individual estrogen development is not affected.

100. (Previously Presented) The improved method of claim 99, wherein the LHRH antagonist is administered by subcutaneous injection.

101. (Canceled)

102. (Previously Presented) The improved method of claim 99, wherein the LHRH antagonist is administered starting cycle day 4 to 8.

103. (Previously Presented) The improved method of claim 99, wherein a daily dose of the LHRH antagonist is administered for 3 to 14 days.

Inventor(s): BOUCHARD *et al.*
Application No.: 08/786,937
Attorney Docket No.: 098501-0235299

104. (Previously Presented) The improved method of claim 99, wherein a daily dose of the LHRH antagonist is administered for 3 to 7 days.

105. (Previously Presented) The improved method of claim 99, wherein ovulation occurs normally, without the administration of a hormone or hormone agonist to induce ovulation.

106. (Previously Presented) The improved method of claim 99, wherein ovulation is induced by administering a hormone or hormone agonist selected from the group consisting of native LH, recombinant LH, an LHRH agonist, and HCG.

107. (Previously Presented) The improved method of claim 99, wherein the LHRH antagonist is Cetrorelix.

108. (Currently Amended) The improved method of claim 99, comprising:
(a) administering human menopausal gonadotropin (HMG) to induce follicle growth, and

(b) administering Cetrorelix to prevent a premature LH surge;
wherein the improvement comprises subcutaneously administering Cetrorelix in a dosage regimen of daily doses of 0.25 mg per day for multiple days;
whereby wherein follicular growth occurs in the absence of a LH surge and a fertilizable oocyte is produced, ovulation occurs between day 9 and 20 of the menstruation cycle, and the Cetrorelix is sufficient to suppress LH, while FSH secretion is maintained at a natural level and individual estrogen development is not affected.

109. (Canceled)

110. (Previously Presented) The improved method of claim 108, wherein Cetrorelix is administered starting cycle day 4 to 8.

111. (Previously Presented) The improved method of claim 108, wherein a daily dose of Cetrorelix is administered for 3 to 14 days.

112. (Previously Presented) The improved method of claim 108, wherein a daily dose of Cetrorelix is administered for 3 to 7 days.

Inventor(s): BOUCHARD *et al.*
Application No.: 08/786,937
Attorney Docket No.: 098501-0235299

113. (Previously Presented) The improved method of claim 108, wherein ovulation occurs normally, without the administration of a hormone or hormone agonist to induce ovulation.

114. (Previously Presented) The improved method of claim 108, wherein ovulation is induced by administering a hormone or hormone agonist selected from the group consisting of native LH, and recombinant LH.

115. (Currently Amended) A method for obtaining the production of a fertilizable oocyte within a program of assisted reproduction techniques comprising:

(a) allowing normal follicular growth and development to proceed in the absence of stimulation by an exogenous gonadotropin;

(b) administering a luteinizing hormone releasing hormone (LHRH) antagonist in a single or dual dosage regimen that prevents a premature LH surge;

~~whereby~~ wherein follicular growth and development proceeds in the absence of a LH surge and a fertilizable oocyte is produced, ovulation occurs between day 9 and 20 of the menstruation cycle, and the ~~Cetrorelix~~ LHRH antagonist is sufficient to suppress LH, while FSH secretion is maintained at a natural level and individual estrogen development is not affected.

116. (Previously Presented) The method of claim 115, wherein the LHRH antagonist is administered by subcutaneous injection.

117. (Canceled)

118. (Previously Presented) The method of claim 115, wherein the LHRH antagonist is administered starting cycle day 4 to 8.

119. (Previously Presented) The method of claim 115, wherein the LHRH antagonist is administered starting on cycle day 6 to 10 and ovulation occurs between day 9 to 16 of the menstruation cycle.

120. (Previously Presented) The method of claim 115, wherein ovulation occurs within 6.5 days following administration of a single or second dose of the LHRH antagonist.

121. (Previously Presented) The method of claim 115, wherein ovulation occurs normally, without the administration of a hormone or hormone agonist to induce ovulation.

Inventor(s): BOUCHARD *et al.*
Application No.: 08/786,937
Attorney Docket No.: 098501-0235299

122. (Previously Presented) The method of claim 115, wherein ovulation is induced by administering a hormone or hormone agonist in order to induce ovulation.

123. (Previously Presented) The method of claim 115, wherein ovulation is induced by administering a hormone or hormone agonist selected from the group consisting of native LH, recombinant LH, an LHRH agonist, and HCG.

124. (Previously Presented) The method of claim 115, wherein the LHRH antagonist is selected from the group consisting of Ganirelix, Antarelix, Antide, Azaline B, Ramorelix, A-76154, Nal-Glu, 88-88, Cetrorelix, a structure-truncated peptide with LHRH-antagonistic activity, a peptidomimetic with LHRH-antagonistic activity, and a bicyclic LHRH-analog with antagonistic activity.

125. (Previously Presented) The method of claim 124, wherein the LHRH antagonist is a peptidomimetic with LHRH-antagonistic activity selected from the group consisting of D-23980 and D-24824.

126. (Previously Presented) The method of claim 124, wherein the LHRH antagonist is Cetrorelix.

127. (Previously Presented) The method of claim 115, wherein a fertilizable oocyte is produced within a program of extracorporeal fertilization by sperm injection.

128. (Previously Presented) The method of claim 115, wherein a fertilizable oocyte is produced within a program of extracorporeal fertilization by *in vitro* fertilization.